REMARKS

This is a preliminary amendment. Claims 1-104 are pending in this application. By this amendment, claims 1-58 are cancelled without prejudice or disclaimer, claims 59-61, 63-64, 66-68, 70-74, 77, 82-83, 88 and 97 are amended, and new claims 105-116 are added. Following entry of this amendment, claims 59-116 will be pending.

Support for the amendments and new claims is found throughout the specification and originally filed claims, including, e.g., at Figures 1, 2A, 20-21, and 23; page 27, lines 34-35; page 3, lines 23-28; page 25, lines 19-29; page 7, lines 29, and 36-37; page 10, lines 17-24; page 11, lines 29-31; page 41, lines 18-20, and 26-27; page 37, line 22; and page 8, line 37 to page 9, line 2. No new matter is added by this amendment. Claim 60 is amended recite "the gene encoding dihydrofolate reductase (DHFR) and the gene encoding glutamine synthetase" (rather than "the genes encoding dihydrofolate reductase (DHFR) and glutamine synthetase"). Claims 61, 66, 88, and 97 are amended to recite "gene encoding DHFR" (rather than "DHFR gene") to improve the consistency of claim language. Compare, eg, claim 60. Claims 61 and 63 are amended to recite "amplifiable second selectable gene" (rather than "amplifiable selectable gene") to improve the consistency of claim language. Compare with claim 59. Claim 63 is also amended to correct a minor typographical error. Claims 64, 74, 77, 82 and 83 are amended to correct claim dependencies. Claim 67 is amended to delete the phrase "comprising a splice donor sequence" and claim 68 is amended and now recite "the intron provides a splicing efficiency of between 80% and 99%", in order to improve the consistency of claim language. Compare, e.g., claim 69. Claim 70 is amended to delete the phrase "wherein the fusion gene is positioned within the intron" in order to delete repetitive language in the claim. Compare claim 67 (which recites this phrase). In addition, the phrase "5" of the intron" is deleted, as this language is implicit in claim 67, from which claim 70 depends. Claim 71 is amended to delete the phrase "wherein the selected sequence and fusion gene are operably linked to the promoter 5' of the selected sequence", in order to delete repetitive language in the claim. Compare claim 67 (which recites this phrase). Claim 72 is rewritten in independent form.

With respect to all amendments and cancelled claims, Applicants have not dedicated or abandoned any unclaimed subject matter and moreover have not acquiesced to any objection and/or rejection made by the Office. Applicants expressly reserve the right to pursue prosecution of any subject matter not presently claimed in one or more future or pending continuation and/or divisional applications.

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Specification

The specification is amended to insert SEQ ID NOs, and to correct minor typographical errors in three citations. The specification is further amended to replace "Figure 9" with "Figure 1" at page 25, lines 21 and 23; page 27, line 35; and page 28, lines 3, 21, and 31. Applicants submit that these references to "Figure 9" are obvious typographical errors for at least the following reasons. First, the paragraphs in which the correction has been made refer to "structures" and "configurations" of exemplary polynucleotide constructs. Figure 1 shows 9 exemplary construct designs. By contrast, Figure 9 depicts graphs showing "DNase productivity vs. GFP productivity" and "DNase RNA vs. DNase productivity". Second, the corrected sentences in the specification refer to line numbers within the figure. Figure 1 contains 9 numbered lines within the figure. By contrast, Figure 9 shows two graphs and does not have any numbered lines. Thus it is evident that one of ordinary skill would understand that these references to Figure 9 are obvious typographic errors, and that Figure 1 should be referenced instead. Accordingly, no new matter is added by the amendments, and entry of the amendments is respectfully requested.

Drawings

The attached sheets of drawing include changes to Figures 20B, 24B, and 24C. Specifically, in Figure 20B, "1920" is changed to "1872"; "1921 to 3381" is changed to 1873 to 3322"; "4200" is changed to "4157"; and "4217 to 4919" is changed to "4158 to 4860". In Figure 24B, "2280" is changed to "2227", and "2287 to 3736" is changed to "2228 to 3677". In Figure 24C, "4800" is changed to "4771" and "4831 to 5533" is changed to "4772 to 5474". These changes correct obvious typographical errors for at least the following reasons. Review of Figures 20 and 24 indicates that the numerals in the column to the left of the polynucleotide sequence refer to the number or position of the last nucleotide in the row associated with the numeral. For example, the numeral "60" at the left of the first line of the polynucleotide sequence in Figure 20A refers to the number or position of the "T" residue at the end of that line. Similarly, the "120" at the left of the second line of the polynucleotide sequence refers to the number or position of the "T" residue at the end of that line, and so on. Turning to Figure 20B, it is evident that the numeral at the left of line 8 of the polynucleotide sequence should be "1872", since the number or position of the last nucleotide in the row, a "C" residue, is 1872. Similarly, the range of numbers defining the insertion site depicted in that row should be "1873 to 3322". Likewise, the numeral at the left of the last line of polynucleotide sequence should be "4157", since the number or position of the last nucleotide in the row (a "C" residue) is 4157, and the range of numbers defining the insertion site depicted in that row should be "4158 to 4860". Similar principles apply to Figure 24B and 24C. Thus it

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is evident that the numerals at the left of line 14 of the polynucleotide sequence of Figure 24B and at the left of line 9 of the polynucleotide sequence of Figure 24C is "2227" and "4771", respectively, and the range of numbers defining the insertion site depicted in those rows should be "2228 to 3677" and "4772 to 5474", respectively. No new matter is added by the changes, and entry of the corrected Figures is respectfully requested.

The Examiner is invited to contact the undersigned in order to expedite the resolution of any remaining issues.

Respectfully submitted, GENENTECH, INC.

Date: 10(13)04

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Amendments to the Drawings:

The attached sheets of drawing include changes to Figures 20B, 24B, and 24C. These sheets replace the original sheets containing Figures 20B, 24B, and 24C. In Figure 20B, "1920" is changed to "1872"; "1921 to 3381" is changed to 1873 to 3322"; "4200" is changed to "4157"; and "4217 to 4919" is changed to "4158 to 4860". In Figure 24B, "2280" is changed to "2227", and "2287 to 3736" is changed to "2228 to 3677". In Figure 24C, "4800" is changed to "4771" and "4831 to 5533" is changed to "4772 to 5474".

Attachments:

Replacement Sheets

Annotated Sheets Showing Changes

In re Application of: Vanessa Chisholm et al. Serial No. 10/714,000

Filed:

November 14, 2003

Title: EXPRESSION VECTORS AND METHODS

APPENDIX

REPLACEMENT FIGURES AND ANNOTATED SHEETS SHOWING CHANGES

Annotated Sheet showing changes

Figure 20B



1500	TTACCAGGAA GCC	ATGAATC AACCAGGCC	CA CCTTAGACTC	TTTGTGACAA	GGATCATGCA
1560	GGAATTTGAA AGT	GACACGT TTTTCCCAG	A AATTGATTTG	GGGAAATATA	AACCTCTCCC
1620	AGAATACCCA GGC	GTCCTCT CTGAGGTCC	A GGAGGAAAA	GGCATCAAGT	ATAAGTTTGA
1680	AGTCTACGAG AAG	AAAGACT AACGTTAAC	T GCTCCCCTCC	TAAAGCTATG	CATTTTTATA
1740	AGACCATGGG ACT	TTTGCTG GCTTTAGAT	C CCCTTGGCTT	CGTTAGAACG	CAGCTACAAT
1800	TAATACATAA CCT	TATGTAT CATACACAT	A CGATTTAGGT	GACACTATAG	ATAACATCCA
1860	CTTTGCCTTT CTC	TCCACAG GTGTCCACT	C CCAGGTCCAA	CTGCACCTCG	GTTCTATCGA
-1920 1872	TTGAATTCCA CC heterologous p	from 1921 to 338 1873 332		site for a	selected
3382	_	GCCC AACTTGTTTA	ጥጥሮሮ እርር መመል		
3420	,	0			
		FAAAGCA ATAGCATCA			·
3480		GGTTTGT CCAAACTCA	-		
3540	TTAATTCGGC GCA	GCACCAT GGCCTGAAA	r aacctctgaa	AGAGGAACTT	GGTTAGGTAC
3600	CTTCTGAGGC GGA	AGAACC AGCTGTGGA	A TGTGTGTCAG	TTAGGGTGTG	GAAAGTCCCC
3660	AGGCTCCCCA GCAC	GCAGAA GTATGCAAA	G CATGCATCTC	AATTAGTCAG	CAACCAGGTG
3720	TGGAAAGTCC CCAG	GGCTCCC CAGCAGGCAC	G AAGTATGCAA	AGCATGCATC	TCAATTAGTC
3780 -	AGCAACCATA GTCC	CGCCCC TAACTCCGCC	CATCCCGCCC	CTAACTCCGC	CCAGTTCCGC
3840	CCATTCTCCG CCCC	ATGGCT GACTAATTTI	TTTTATTTAT	GCAGAGGCCG	AGGCCGCCTC
3900	GGCCTCTGAG CTAT	TCCAGA AGTAGTGAGG	G AGGCTTTTTT	GGAGGAGCTT	TTGCAAAAAG
3960	CTAGCTTATC CGGC	CGGGAA CGGTGCATTG	GAACGCGGAT	TCCCCGTGCC	AAGAGTCAGG
4020	TAAGTACCGC CTAT	AGAGTC TATAGGCCCA	CCCCCTTGGC	TTCGTTAGAA (CGCGGCTACA
4080	ATTAATACAT AACC	TTTTGG ATCGATCCTA	CTGACACTGA	CATCCACTTT	TTCTTTTTCT
4140	CCACAGGTGT CCAC	TCCCAG GTCCAACTGC	ACCTCGGTTC	GCGAAGCTAG (CTTGGGCTGC
4 20 0 4157	ATCGATTGAA TTCC	ACC <from 4<del="">217 to 415%</from>	4 919 , inser 4 <i>860</i>	tion site fo	or a
	selected hetero	logous polypeptid			

Annotated Sheet showing changes

Figure 24B

1500	ACCATTGAAC	TGCATCGTCG	CCGTGTCCCA	AAATATGGGG	ATTGGCAAGA	ACGGAGACC'
1560	ACCCTGGCCT	CCGCTCAGGA	ACGCGTTCAA	GTACTTCCAA	AGAATGACCA	CAACCTCTTC
1620	AGTGGAAGGT	AAACAGAATC	TGGTGATTAT	GGGTAGGAAA	ACCTGGTTCT	CCATTCCTG
1680	GAAGAATCGA	CCTTTAAAGG	ACAGAATTAA	TATAGTTCTC	AGTAGAGAAC	TCAAAGAACO
1740	ACCACGAGGA	GCTCATTTTC	TTGCCAAAAG	TTTGGATGAT	GCCTTAAGAC	TTATTGAACA
1800	ACCGGAATTG	GCAAGTAAAG	TAGACATGGT	TTGGATAGTC	GGAGGCAGTT	CTGTTTACCA
1860	GGAAGCCATG	AATCAACCAG	GCCACCTCAG	ACTCTTTGTG	ACAAGGATCA	TGCAGGAATI
1920	TGAAAGTGAC	ACGTTTTTCC	CAGAAATTGA	TTTGGGGAAA	TATAAACCTC	TCCCAGAATA
1980	CCCAGGCGTC	CTCTCTGAGG	TCCAGGAGGA	AAAAGGCATC	AAGTATAAGT	TTGAAGTCTA
2040	CGAGAAGAAA	GACTAACGTT	AACTGCTCCC	CTCCTAAAGC	TATGCATTTT	TATAAGACCA
2100	TGAGACTTTT	GCTGGCTTTA	GATCCCCTTG	GCTTCGTTAG	AACGCAGCTA	CAATTAATAC
2160	ATAACCTTAT	GTATCATACA	CATACGATTT	AGGTGACACT	ATAGAATAAC	ATCCACTTTG
2220	CCTTTCTCTC	CACAGGTGTC	CACTCCCAGG	TCCAACTGCA	CCTCGGTTCT	ATCGATTGAA
2280 222フ	TTCCACC <f1< th=""><th>rom 2287 to</th><th>3736, inser</th><th>rtion site</th><th>for a selec</th><th>ted</th></f1<>	rom 2287 to	3 736 , inser	rtion site	for a selec	ted
	heterologou	ıs polypepti				
3737	CGA TGGCCGC	CCAT GGCCCAA	CTT GTTTAT	GCA GCTTATA	AATG	
3780	GTTACAAATA	AAGCAATAGC	ATCACAAATT	TCACAAATAA	AGCATTTTT	TCACTGCATT
3840	CTAGTTGTGG	TTTGTCCAAA	CTCATCAATG	TATCTTATCA	TGTCTGGATC	GGGAATTAAT
3900	TCGGCGCAGC	ACCATGGCCT	GAAATAACCT	CTGAAAGAGG	AACTTGGTTA	GGTACCTATT
3960	AATAGTAATC	AATTACGGGG	TCATTAGTTC	ATAGCCCATA	TATGGAGTTC	CGCGTTACAT
4020	AACTTACGGT	AAATGGCCCG	CCTGGCTGAC	CGCCCAACGA	CCCCGCCCA	TTGACGTCAA
4080	TAATGACGTA	TGTTCCCATA	GTAACGCCAA	TAGGGACTTT	CCATTGACGT	CAATGGGTGG
4140	AGTATTTACG	GTAAACTGCC	CACTTGGCAG	TACATCAAGT	GTATCATATG	CCAAGTACGC
4200	CCCCTATTGA	CGTCAATGAC	GGTAAATGGC	CCGCCTGGCA	TTATGCCCAG	TACATGACCT
4260	TATGGGACTT	TCCTACTTGG	CAGTACATCT	ACGTATTAGT	CATCGCTATT	ACCATGGTGA



Annotated Sheet showing changes

Figure 24C

4320	TGCGGTTTTG GCAGTACATC AATGGGCGTG GATAGCGGTT TGACTCACGG GGATTTCCA
4380	GTCTCCACCC CATTGACGTC AATGGGAGTT TGTTTTGGCA CCAAAATCAA CGGGACTTTC
4440	CAAAATGTCG TAACAACTCC GCCCCATTGA CGCAAATGGG CGGTAGGCGT GTACGGTGGC
4500	AGGTCTATAT AAGCAGAGCT CGTTTAGTGA ACCGTCAGAT CGCCTGGAGA CGCCATCCAC
4560	GCTGTTTTGA CCTGCTAGCT TATCCGGCCG GGAACGGTGC ATTGGAACGC GGATTCCCCG
4620	TGCCAAGAGT CAGGTAAGTA CCGCCTATAG AGTCTATAGG CCCACCCCCT TGGCTTCGTT
4680	AGAACGCGGC TACAATTAAT ACATAACCTT TTGGATCGAT CCTACTGACA CTGACATCCA
4740	CTTTTTCTTT TTCTCCACAG GTGTCCACTC CCAGGTCCAA CTGCACCTCG GTTCGCGAAG
4800	CTCGCTTGGG CTGCATCGAT TGAATTCCAC C <from 4831="" 5533,="" insertion<="" th="" to=""></from>
4771	site for a selected heterologous polypeptide>
5534	CGATGG CCGCCATGGC CCAACTTGTT TATTGCAGCT TATAATGGTT
5580	ACAAATAAAG CAATAGCATC ACAAATTTCA CAAATAAAGC ATTTTTTTCA CTGCATTCTA
5640	GTTGTGGTTT GTCCAAACTC ATCAATGTAT CTTATCATGT CTGGATCGGG AATTAATTCG
5700	GCGCAGCACC ATGGCCTGAA ATAAGTTTAA ACCCTCTGAA AGAGGAACTT GGTTAGGTAC
5760	CGACTAGTCT TTTGCAAAAA GCTGTTACCT CGAGCGGCCG CTTAATTAAG GCGCGCCATT
5820	TAAATCCTGC AGGTAACAGC TTGGCACTGG CCGTCGTTTT ACAACGTCGT GACTGGGAAA
5880	ACCCTGGCGT TACCCAACTT AATCGCCTTG CAGCACATCC CCCTTTCGCC AGCTGGCGTA
5940	ATAGCGAAGA GGCCCGCACC GATCGCCCTT CCCAACAGTT GCGCAGCCTG AATGGCGAAT
6000	GGCGCCTGAT GCGGTATTTT CTCCTTACGC ATCTGTGCGG TATTTCACAC CGCATACGTC
6060	AAAGCAACCA TAGTACGCGC CCTGTAGCGG CGCATTAAGC GCGGCGGGTG TGGTGGTTAC
6120	GCGCAGCGTG ACCGCTACAC TTGCCAGCGC CCTAGCGCCC GCTCCTTTCG CTTTCTTCCC
6180	TTCCTTTCTC GCCACGTTCG CCGGCTTTCC CCGTCAAGCT CTAAATCGGG GGCTCCCTTT
	AGGGTTCCGA TTTAGTGCTT TACGGCACCT CGACCCCAAA AAACTTGATT TGGGTGATGG
	TTCACGTAGT GGGCCATCGC CCTGATAGAC GGTTTTTCGC CCTTTGACGT TGGAGTCCAC
	GTTCTTTAAT AGTGGACTCT TGTTCCAAAC TGGAACAACA CTCAACCCTA TCTCGGGCTA
	TTCTTTTGAT TTATAAGGGA TTTTGCCGAT TTCGGCCTAT TGGTTAAAAA ATGAGCTGAT